CLAIMS

( all claim pages)

#### We claim:

#### A compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof; wherein R<sub>1</sub> is hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl;

 $R_2$  and  $R_3$  are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, or an aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl moiety; or

R<sub>1</sub> and R<sub>2</sub>, when taken together, may form a substituted or unsubstituted, saturated or unsaturated cyclic ring of 3 to 8 carbon atoms; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a substituted or unsubstituted, saturated or unsaturated cyclic ring of 3 to 8 carbon atoms;

R4 is hydrogen or halogen;

 $\mathbb{R}_{s}$  is hydrogen, an oxygen protecting group or a prodrug;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;

**n** is 0-2;

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or an aliphatic moiety optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $\mathbb{R}_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $INR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2$ - $R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2$ - $R_{14}$ ;



wherein R<sub>12</sub> and R<sub>13</sub> are, independently for each occurrence, hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl; or a protecting group, or R<sub>12</sub> and R<sub>13</sub>, taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of R<sub>12</sub> and R<sub>13</sub> are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or -N(alkyl), or wherein  $X_2$ - $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is  $-(C=O)NHR_{15}$   $-(C=O)OR_{15}$ , or  $-(C=O)R_{15}$ , wherein each occurrence of  $R_{15}$  is independently hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl; or  $R_{14}$  is  $-SO_2(R_{16})$ , wherein  $R_{16}$  is an aliphatic moiety, wherein one or more of  $R_{14}$ ,  $R_{15}$ , or  $R_{16}$  are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

 $R_{10}$  is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;  $R_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH2 or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or aliphatic, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond;

with the proviso that when n is 1; X is O;  $R_1$  is methyl;  $R_2$ ,  $R_3$ ,  $R_7$  and  $R_{11}$  are each hydrogen;  $R_5$  is hydrogen,  $C_{1-4}$ alkyl or  $-C(=O)C_{1-4}$ alkyl;  $R_6$  is hydrogen, OH,  $C_{1-4}$ alkoxy or  $-OC(=O)C_{1-4}$ alkyl; and  $R_9$  is OH,  $C_{1-4}$ alkoxy or  $-OC(=O)C_{1-4}$ alkyl; then one or more if the following groups do not occur simultaneously as defined:



- (i) R<sub>4</sub> is hydrogen; R<sub>10</sub> and R<sub>8</sub> are independently OH, C<sub>1-4</sub>alkoxy or OC(=O)C<sub>1-4</sub>alkyl; and Y-Z is –CH<sub>2</sub>CH<sub>2</sub>- or –CH=CH-;
- (ii) R<sub>4</sub> and R<sub>8</sub> are each hydrogen; R<sub>10</sub> is OH, C<sub>1-4</sub>alkoxy or -OC(=O)C<sub>1-</sub>

4alkyl; and Y-Z is -CHRYCHRZ-, -CH=CH- or ; wherein  $R^Y$  and  $R^Z$  are independently hydrogen,  $C_{1-4}$ alkyl or  $C_{1-4}$ alkanoyl; and

- (iii) R<sub>4</sub> and R<sub>10</sub> are each hydrogen, OH, C<sub>1-4</sub>alkoxy or -OC(=O)C<sub>1-4</sub>alkyl; R<sub>8</sub> is hydrogen, OH, halogen, C<sub>1-4</sub>alkoxy or -OC(=O)C<sub>1-4</sub>alkyl; and Y-Z is -CH<sub>2</sub>CH<sub>2</sub>-, -CH=CH- or -C(=O)CH<sub>2</sub>-.
- 2. The compound of claim 1, where the following groups do not occur simultaneously as defined:

X is oxygen,

R<sub>1</sub> is methyl,

R<sub>2</sub> and R<sub>3</sub> are each hydrogen,

R<sub>4</sub> is hydrogen,

R<sub>5</sub> is hydrogen, C<sub>1-6</sub>alkyl or C<sub>1-6</sub>alkanoyl,

 $R_6$  is OR', where R' is hydrogen,  $C_{1-6}$ alkyl or  $C_{1-6}$ alkanoyl with Sconfiguration,

R<sub>7</sub> is hydrogen,

Y and Z together represent  $-CHR_{17}$ - $CHR_{18}$ -or  $-CR_{17}$ = $CR_{18}$ -, wherein  $R_{17}$  and  $R_{18}$  are independently hydrogen, or when Y and Z are  $-CHR_{17}$ - $CHR_{18}$ ,  $R_{17}$  and  $R_{18}$  taken together are -O-;

 $R_8$  is hydrogen or OR', where R' is hydrogen,  $C_{1-6}$ alkyl or  $C_{1-6}$ alkanoyl,

 $R_9$  is OR', where R' is hydrogen,  $C_{1-6}$ alkyl or  $C_{1-6}$ alkanoyl; and  $R_{10}$  is OR", where R" is hydrogen,  $C_{1-6}$ alkyl or  $C_{1-6}$ alkanoyl; and  $R_{11}$  is hydrogen.

3. The compound of claim 1, wherein:

 $\mathbb{R}_1$  is hydrogen, straight or branched  $\mathbb{C}_{1-6}$ alkyl, straight or branched  $\mathbb{C}_{1-6}$ 6heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl;

R<sub>2</sub> and R<sub>3</sub> are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, straight or branched C<sub>1-6</sub>heteroalkyl, or aryl, wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl; or

R<sub>1</sub> and R<sub>2</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen; or

 $R_1$  and  $R_3$ , when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen;

R4 is hydrogen or halogen;

 $R_5$  is hydrogen or a protecting group;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;

n is 0-2;

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2$ - $R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2$ - $R_{14}$ ;

wherein  $R_{12}$  and  $R_{13}$  are, independently for each occurrence, hydrogen,  $C_{1-6}$  alkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or a protecting group, or  $R_{12}$  and  $R_{13}$ , taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of  $R_{12}$  and  $R_{13}$  are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,



wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or -N(alkyl), or wherein  $X_2$ - $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is  $-(C=O)NHR_{15}$   $-(C=O)OR_{15}$ , or  $-(C=O)R_{15}$ , wherein each occurrence of  $R_{15}$  is independently hydrogen, alkyl, heteroalkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or  $R_{14}$  is  $-SO_2(R_{16})$ , wherein  $R_{16}$  is an alkyl moiety, wherein one or more of  $R_{14}$ ,  $R_{15}$ , or  $R_{16}$  are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

R<sub>10</sub> is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;
R<sub>11</sub> is hydrogen, hydroxyl or protected hydroxyl;
X is absent or is O, NH, N-alkyl, CH<sub>2</sub> or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or C<sub>1-6</sub>alkyl, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond.

- 4. The compound of claim 3, where X is oxygen and n is 1.
- 5. The compound of claim 3, where R<sub>4</sub> is halogen.
- 6. The compound of claim 3, where R<sub>4</sub> is fluorine.
- 7. The compound of claim 3, where Y and Z together represent-CH=CH-
- 8. The compound of claim 3, where Y and Z together represent trans—CH=CH-.



9. The compound of claim 3, wherein  $R_1$  and  $R_2$  are each methyl and  $R_3$  is hydrogen and the compound has the structure:

wherein  $R_4$ - $R_{11}$ , n, X, Y and Z are as defined in claim 3.

- 10. The compound of claim 9, wherein X is oxygen and n is 1.
- 11. The compound of claim 9, wherein R<sub>4</sub> is halogen.

- 20. The compound of claim 15, wherein X is oxygen, n is 1, R<sub>1</sub> and R<sub>2</sub> are each methyl, R<sub>3</sub> is hydrogen, R<sub>4</sub> is halogen, and Y and Z together represent -CH=CH-.
- 21. The compound of claim 18 or 20, wherein -CH=CH- is trans.
- 22. A compound having the structure:

23. A compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

24. A compound having the structure:

#### 25. A compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

# 26. A compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

# 27. A compound having the structure:



### 28. A compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

# 29. A compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

### 30. A compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

#### 31. A compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

#### 32. A compound having the structure:

### 33. A compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

# 34. A compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

### 35. A compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

# 36. A compound having the structure:

# 37. A pharmaceutical composition comprising: a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof; wherein  $\mathbf{R}_{\mathbf{I}}$  is hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl;

R<sub>2</sub> and R<sub>3</sub> are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, or an aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl moiety; or

 $R_1$  and  $R_2$ , when taken together, may form a substituted or unsubstituted, saturated or unsaturated cyclic ring of 3 to 8 carbon atoms; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a substituted or unsubstituted, saturated or unsaturated cyclic ring of 3 to 8 carbon atoms;

R4 is hydrogen or halogen;

R<sub>5</sub> is hydrogen, an oxygen protecting group or a prodrug;

 $\mathbb{R}_6$  is hydrogen, hydroxyl, or protected hydroxyl;

n is 0-2;



 $\mathbb{R}_{7}$ , for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or an aliphatic moiety optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2$ - $R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2$ - $R_{14}$ ;

wherein R<sub>12</sub> and R<sub>13</sub> are, independently for each occurrence, hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl; or a protecting group, or R<sub>12</sub> and R<sub>13</sub>, taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of R<sub>12</sub> and R<sub>13</sub> are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or -N(alkyl), or wherein  $X_2$ - $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is – (C=O)NHR<sub>15</sub> –(C=O)OR<sub>15</sub>, or –(C=O)R<sub>15</sub>, wherein each occurrence of  $R_{15}$  is independently hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl; or  $R_{14}$  is –SO<sub>2</sub>( $R_{16}$ ), wherein  $R_{16}$  is an aliphatic moiety, wherein one or more of  $R_{14}$ ,  $R_{15}$ , or  $R_{16}$  are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

R<sub>10</sub> is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino; R<sub>11</sub> is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH2 or S;



Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or aliphatic, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond; and

a pharmaceutically acceptable carrier;

with the proviso that when n is 1; X is O;  $R_1$  is methyl;  $R_2$ ,  $R_3$ ,  $R_7$  and  $R_{11}$  are each hydrogen;  $R_5$  is hydrogen,  $C_{1-4}$ alkyl or  $-C(=0)C_{1-4}$ alkyl;  $R_6$  is hydrogen, OH,  $C_{1-4}$ alkoxy or  $-OC(=0)C_{1-4}$ alkyl; and  $R_9$  is OH,  $C_{1-4}$ alkoxy or  $-OC(=0)C_{1-4}$ alkyl; then one or more if the following groups do not occur simultaneously as defined:

- (i) R<sub>4</sub> is hydrogen; R<sub>10</sub> and R<sub>8</sub> are independently OH, C<sub>1-4</sub>alkoxy or OC(=0)C<sub>1-4</sub>alkyl; and Y-Z is –CH<sub>2</sub>CH<sub>2</sub>- or –CH=CH-;
- (ii)  $R_4$  and  $R_8$  are each hydrogen;  $R_{10}$  is OH,  $C_{1-4}$ alkoxy or  $-OC(=O)C_{1-4}$ .

4alkyl; and Y-Z is -CHRYCHRZ-, -CH=CH- or ', wherein RY and RZ are independently hydrogen, C<sub>1-4</sub>alkyl or C<sub>1-4</sub>alkanoyl; and

- (iii) R<sub>4</sub> and R<sub>10</sub> are each hydrogen, OH, C<sub>1-4</sub>alkoxy or -OC(=O)C<sub>1-4</sub>alkyl; R<sub>8</sub> is hydrogen, OH, halogen, C<sub>1-4</sub>alkoxy or -OC(=O)C<sub>1-4</sub>alkyl; and Y-Z is -CH<sub>2</sub>CH<sub>2</sub>-, -CH=CH- or -C(=O)CH<sub>2</sub>-.
- 38. The pharmaceutical composition of claim 37, wherein the compound is present in an amount effective to inhibit NF-kB activation.
- 39. The pharmaceutical composition of claim 37, wherein the compound is present in an amount effective to inhibit AP-1 activation.
- 40. The pharmaceutical composition of claim 37, wherein the compound is present in an amount effective to inhibit a protein kinase.
- 41. The pharmaceutical composition of claim 39, wherein the protein kinase is MEKK1, MEK1, VEGFr or PDGFr.



- 42. The pharmaceutical composition of claim 37, wherein the compound is present in an amount effective to inhibit proliferation of cancerous cells and angiogenesis on solid tumors.
- 43. The pharmaceutical composition of claim 37, wherein the compound is present in an amount effective to have an anti-inflammatory effect.
- 44. The pharmaceutical composition of claim 37, wherein the compound is present in an amount effective to treat psoriasis.
- 45. The pharmaceutical composition of claim 37, wherein the compound is present in an amount effective to reduce skin photodamage.
- 46. The pharmaceutical composition of claim 37, wherein the compound is present in an amount effective to prevent restensis.
- 47. The pharmaceutical composition of claim 37, where:

 $\mathbf{R}_1$  is hydrogen, straight or branched  $C_{1\text{-}6}$ alkyl, straight or branched  $C_{1\text{-}6}$ heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl;

R<sub>2</sub> and R<sub>3</sub> are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, straight or branched C<sub>1-6</sub>heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl; or

 $R_1$  and  $R_2$ , when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen;



R<sub>4</sub> is hydrogen or halogen;
R<sub>5</sub> is hydrogen or a protecting group;
R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;
n is 0-2;

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2$ - $R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2$ - $R_{14}$ ;

wherein  $R_{12}$  and  $R_{13}$  are, independently for each occurrence, hydrogen,  $C_{1-6}$ alkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or a protecting group, or  $R_{12}$  and  $R_{13}$ , taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of  $R_{12}$  and  $R_{13}$  are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or -N(alkyl), or wherein  $X_2$ - $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is  $-(C=O)NHR_{15}$   $-(C=O)OR_{15}$ , or  $-(C=O)R_{15}$ , wherein each occurrence of  $R_{15}$  is independently hydrogen, alkyl, heteroalkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or  $R_{14}$  is  $-SO_2(R_{16})$ , wherein  $R_{16}$  is an alkyl moiety, wherein one or more of  $R_{14}$ ,  $R_{15}$ , or  $R_{16}$  are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;



 $\mathbb{R}_{10}$  is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;  $\mathbb{R}_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH2 or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or C<sub>1-6</sub>alkyl, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond.

- 48. The pharmaceutical composition of claim 47, where X is oxygen and n is 1.
- 49. The pharmaceutical composition of claim 47, where R<sub>4</sub> is halogen.
- 50. The pharmaceutical composition of claim 49, where R<sub>4</sub> is fluorine.
- 51. The pharmaceutical composition of claim 47, where Y and Z together represent CH=CH-.
- 52. The pharmaceutical composition of claim 51, wherein -CH=CH- is trans.

wherein R<sub>1</sub>-R<sub>13</sub>, n, X, Y and Z are as defined in claim 46, or

R<sub>13</sub> and R<sub>8</sub> may, when taken together, form a cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydrogen, alkyloxy, amino, alkylamino, aminoalkyl, and halogen.

- 60. The pharmaceutical composition of claim 59, wherein X is oxygen and n is 1.
- 61. The pharmaceutical composition of claim 59, wherein R<sub>4</sub> is halogen.
- 62. The pharmaceutical composition of claim 59, wherein Y and Z together represent CH=CH-.
- 63. The pharmaceutical composition of claim 59, wherein  $R_1$  and  $R_2$  are each methyl and  $R_3$  is hydrogen.
- 64. The pharmaceutical composition of claim 59 wherein X is oxygen, n is 1, R<sub>1</sub> and R<sub>2</sub> are each methyl, R<sub>3</sub> is hydrogen, R<sub>4</sub> is halogen, and Y and Z together represent -CH=CH-.
- 65. The pharmaceutical composition of claim 63 or 64 wherein -CH=CH- is trans.
- 66. A pharmaceutical composition comprising: a compound having the structure:

67. A pharmaceutical composition comprising: a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof; and a pharmaceutically acceptable carrier.

69. A pharmaceutical composition comprising: a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof; and a pharmaceutically acceptable carrier.

70. A pharmaceutical composition comprising: a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof; and a pharmaceutically acceptable carrier.

72. A pharmaceutical composition comprising: a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof; and a pharmaceutically acceptable carrier.

74. A pharmaceutical composition comprising: a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof; and a pharmaceutically acceptable carrier.

75. A pharmaceutical composition comprising: a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof, and a pharmaceutically acceptable carrier.

77. A pharmaceutical composition comprising: a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof; and a pharmaceutically acceptable carrier.

78. A pharmaceutical composition comprising: a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof; and a pharmaceutically acceptable carrier.

79. A pharmaceutical composition comprising:



a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof; and a pharmaceutically acceptable carrier.

80. A pharmaceutical composition comprising: a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof; and a pharmaceutically acceptable carrier.

81. A topical pharmaceutical composition for preventing or treating UVB-induced photodamage comprising:

a compound having the structure:

$$R_{11}$$
 $R_{10}$ 
 $R_{11}$ 
 $R_{11}$ 
 $R_{11}$ 
 $R_{11}$ 
 $R_{11}$ 
 $R_{11}$ 
 $R_{12}$ 
 $R_{13}$ 
 $R_{14}$ 
 $R_{15}$ 
 $R$ 

or pharmaceutically acceptable salt, ester, or salt of ester thereof; wherein  $R_1$  is hydrogen, straight or branched  $C_{1-6}$ alkyl, straight or branched  $C_{1-6}$ heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl;

R<sub>2</sub> and R<sub>3</sub> are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, straight or branched C<sub>1-6</sub>heteroalkyl, or aryl, wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl; or

 $R_1$  and  $R_2$ , when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen;

R4 is hydrogen or halogen;

R<sub>5</sub> is hydrogen or a protecting group;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl:

m is 0-2:

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2$ - $R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2$ - $R_{14}$ ;

wherein  $R_{12}$  and  $R_{13}$  are, independently for each occurrence, hydrogen,  $C_{1\text{-}6}$ alkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or a protecting group, or  $R_{12}$  and  $R_{13}$ , taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of  $R_{12}$  and  $R_{13}$  are optionally further substituted with one or more occurrences



of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or -N(alkyl), or wherein  $X_2$ - $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is  $-(C=O)NHR_{15}$   $-(C=O)OR_{15}$ , or  $-(C=O)R_{15}$ , wherein each occurrence of  $R_{15}$  is independently hydrogen, alkyl, heteroalkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or  $R_{14}$  is  $-SO_2(R_{16})$ , wherein  $R_{16}$  is an alkyl moiety, wherein one or more of  $R_{14}$ ,  $R_{15}$ , or  $R_{16}$  are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

 $R_{10}$  is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;  $R_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH<sub>2</sub> or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or C<sub>1-6</sub>alkyl, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond; and

a pharmaceutically acceptable carrier;

wherein the compound is present in an amount effective to prevent or treat UVB-induced photodamage.

- 82. The pharmaceutical composition of claim 81, further comprising a cosmetic ingredient.
- 83. The pharmaceutical composition of claim 82, wherein the cosmetic ingredient is a sunscreen.



84. A method for treating an inflammatory and/or autoimmune disorder or a disorder resulting from increased angiogenesis and/or cell proliferation comprising:

administering to a subject in need thereof a therapeutically effective amount of a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof; wherein  $R_1$  is hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl;

R<sub>2</sub> and R<sub>3</sub> are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, or an aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl moiety; or

 $R_1$  and  $R_2$ , when taken together, may form a substituted or unsubstituted, saturated or unsaturated cyclic ring of 3 to 8 carbon atoms; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a substituted or unsubstituted, saturated or unsaturated cyclic ring of 3 to 8 carbon atoms;

R<sub>4</sub> is hydrogen or halogen;

R<sub>5</sub> is hydrogen, an oxygen protecting group or a prodrug;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;

n is 0-2;

 $\mathbb{R}_{7}$ , for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or an aliphatic moiety optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;



 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2$ - $R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2$ - $R_{14}$ ;

wherein R<sub>12</sub> and R<sub>13</sub> are, independently for each occurrence, hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl; or a protecting group, or R<sub>12</sub> and R<sub>13</sub>, taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of R<sub>12</sub> and R<sub>13</sub> are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or -N(alkyl), or wherein  $X_2$ - $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is - (C=O)NHR<sub>15</sub> - (C=O)OR<sub>15</sub>, or - (C=O)R<sub>15</sub>, wherein each occurrence of  $R_{15}$  is independently hydrogen, aliphatic, heteroaliphatic, alicyclic, heteroalicyclic, aryl or heteroaryl; or  $R_{14}$  is -SO<sub>2</sub>( $R_{16}$ ), wherein  $R_{16}$  is an aliphatic moiety, wherein one or more of  $R_{14}$ ,  $R_{15}$ , or  $R_{16}$  are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

 $R_{10}$  is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;  $R_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH2 or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or aliphatic, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond; and

a pharmaceutically acceptable carrier or diluent;



with the proviso that when n is 1; X is O;  $R_1$  is methyl;  $R_2$ ,  $R_3$ ,  $R_7$  and  $R_{11}$  are each hydrogen;  $R_5$  is hydrogen,  $C_{1-4}$ alkyl or  $-C(=O)C_{1-4}$ alkyl;  $R_6$  is hydrogen, OH,  $C_{1-4}$ alkoxy or  $-OC(=O)C_{1-4}$ alkyl; and  $R_9$  is OH,  $C_{1-4}$ alkoxy or  $-OC(=O)C_{1-4}$ alkyl; then one or more if the following groups do not occur simultaneously as defined:

- (i) R<sub>4</sub> is hydrogen; R<sub>10</sub> and R<sub>8</sub> are independently OH, C<sub>1-4</sub>alkoxy or OC(=O)C<sub>1-4</sub>alkyl; and Y-Z is -CH<sub>2</sub>CH<sub>2</sub>- or -CH=CH-; and
- (ii) R<sub>4</sub> and R<sub>8</sub> are each hydrogen; R<sub>10</sub> is OH, C<sub>1-4</sub>alkoxy or -OC(=O)C<sub>1</sub>.

  4alkyl; and Y-Z is -CHR<sup>Y</sup>CHR<sup>Z</sup>-, -CH=CH- or ; wherein R<sup>Y</sup> and R<sup>Z</sup> are independently hydrogen, C<sub>1-4</sub>alkyl or C<sub>1-4</sub>alkanoyl; and
- (iii) R<sub>4</sub> and R<sub>10</sub> are each hydrogen, OH, C<sub>1-4</sub>alkoxy or -OC(=O)C<sub>1-4</sub>alkyl; R<sub>8</sub> is hydrogen, OH, halogen, C<sub>1-4</sub>alkoxy or -OC(=O)C<sub>1-4</sub>alkyl; and Y-Z is -CH<sub>2</sub>CH<sub>2</sub>-, -CH=CH- or -C(=O)CH<sub>2</sub>-; whereby the compound induces mRNA degradation and the method is for treating a disorder resulting from cell proliferation.
- 85. The method of claim 84, wherein the method is for treating a disorder selected from the group consisting of rheumatoid arthritis, psoriasis, asthma, cancer, sepsis, inflammatory bowel disease, atopic dermatitis, Crohn's disease, and autoimmune disorders.
- 86. The method of claim 84, wherein the method is for treating rheumatoid arthritis.
- 87. The method of claim 84, wherein the method is for treating psoriasis.
- 88. The method of claim 84, wherein the method is for treating asthma.
- 89. The method of claim 84, wherein:

  Rais hydrogen straight or branched Casally attrait

 $R_1$  is hydrogen, straight or branched  $C_{1\text{-}6}$ alkyl, straight or branched  $C_{1\text{-}6}$ heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl;

 $R_2$  and  $R_3$  are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, straight or branched  $C_{1-6}$ alkyl, straight or branched  $C_{1-6}$ heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl; or

R<sub>1</sub> and R<sub>2</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen;

R4 is hydrogen or halogen;

R<sub>5</sub> is hydrogen or a protecting group;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;

n is 0-2;

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2$ - $R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2$ - $R_{14}$ ;

wherein  $R_{12}$  and  $R_{13}$  are, independently for each occurrence, hydrogen,  $C_{1-6}$ alkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or a protecting group, or  $R_{12}$  and  $R_{13}$ , taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of  $R_{12}$  and  $R_{13}$  are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or -N(alkyl), or wherein  $X_2$ - $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is  $-(C=0)NHR_{15}$   $-(C=0)OR_{15}$ , or  $-(C=0)R_{15}$ , wherein each occurrence of  $R_{15}$  is independently hydrogen, alkyl, heteroalkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or  $R_{14}$  is  $-SO_2(R_{16})$ , wherein  $R_{16}$  is an alkyl moiety, wherein one or more of  $R_{14}$ ,  $R_{15}$ , or  $R_{16}$  are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

 $R_{10}$  is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;  $R_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH2 or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or C<sub>1-6</sub>alkyl, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond.

- 90. The method of claim 89, wherein in the compound X is oxygen and n is 1.
- 91. The method of claim 89, wherein in the compound R<sub>4</sub> is halogen.
- 92. The method of claim 89 is wherein in the compound R<sub>4</sub> is fluorine.
- 93. The method of claim 89, wherein in the compound Y and Z together represent-CH=CH-



- 94. The method of claim 93, wherein in the compound Y and Z together represent trans -CH=CH-.
- 95. The method of claim 89, comprising administering a compound wherein  $R_1$  and  $R_2$  are each methyl and  $R_3$  is hydrogen and the compound has the structure:

wherein R<sub>4</sub>-R<sub>11</sub>, n, X, Y and Z are as defined in claim 88.

- 96. The method of claim 95, wherein in the compound X is oxygen and n is 1.
- 97. The method of claim 95, wherein in the compound R<sub>4</sub> is halogen.

- 106. The method of claim 101, wherein in the compound X is oxygen, n is 1,  $R_1$  and  $R_2$  are each methyl,  $R_3$  is hydrogen,  $R_4$  is halogen, and Y and Z together represent -CH=CH-.
- 107. The method of claim 105 or 106, wherein in the compound -CH=CH- is trans.
- 108. The method of claim 84, comprising administering a compound having the structure:

109. The method of claim 84, comprising administering a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

110. The method of claim 84, comprising administering a compound having the structure:



111. The method of claim 84, comprising administering a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

112. The method of claim 84, comprising administering a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

113. The method of claim 84, comprising administering a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

114. The method of claim 84, comprising administering a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

115. The method of claim 84, comprising administering a compound having the structure:

116. The method of claim 84, comprising administering a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

117. The method of claim 84, comprising administering a compound having the structure:

or pharmaceutically acceptable salt, ester, or salt of ester thereof.

118. The method of claim 84, comprising administering a compound having the structure:

119. A method for providing protection against UVB-induced photodamage to a subject, said method comprising:

Administering to the subject in need thereof a composition comprising a compound having the structure:

$$R_{11}$$
 $R_{10}$ 
 $R_{11}$ 
 $R$ 

or pharmaceutically acceptable salt, ester, or salt of ester thereof; wherein  $\mathbb{R}_1$  is hydrogen, straight or branched  $C_{1\text{-}6}$ alkyl, straight or branched  $C_{1\text{-}6}$ heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl;

 $\mathbb{R}_2$  and  $\mathbb{R}_3$  are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, straight or branched  $C_{1\text{-}6}$ alkyl, straight or branched  $C_{1\text{-}6}$ heteroalkyl, or aryl,



wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl; or

 $R_1$  and  $R_2$ , when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen;

R<sub>4</sub> is hydrogen or halogen;

R<sub>5</sub> is hydrogen or a protecting group;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;

n is 0-2;

 $R_7$ , for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2$ - $R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2$ - $R_{14}$ ;

wherein  $R_{12}$  and  $R_{13}$  are, independently for each occurrence, hydrogen,  $C_{1\text{-}6}$ alkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or a protecting group, or  $R_{12}$  and  $R_{13}$ , taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of  $R_{12}$  and  $R_{13}$  are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or -N(alkyl), or wherein  $X_2$ - $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is – (C=O)NHR<sub>15</sub> –(C=O)OR<sub>15</sub>, or –(C=O)R<sub>15</sub>, wherein each



occurrence of  $R_{15}$  is independently hydrogen, alkyl, heteroalkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or  $R_{14}$  is  $-SO_2(R_{16})$ , wherein  $R_{16}$  is an alkyl moiety, wherein one or more of  $R_{14}$ ,  $R_{15}$ , or  $R_{16}$  are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

 $R_{10}$  is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;  $R_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH2 or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of  $R_{17}$  and  $R_{18}$  is independently hydrogen or  $C_{1-6}$ alkyl, or  $R_{17}$  and  $R_{18}$  taken together is -O-, -CH<sub>2</sub>- or -NR<sub>19</sub>-, wherein  $R_{19}$  is hydrogen or  $C_{1-6}$ alkyl, and Y and Z may be connected by a single or double bond; and

a pharmaceutically acceptable carrier or diluent.

- 120. The method of claim 119, wherein in the step of administering, the composition is administered topically.
- 121. The method of claim 119, wherein the photodamage is skin wrinkles.
- 122. The method of claim 119, wherein the photodamage is a skin cancer.
- 123. A method for preventing or reducing the rate of restenosis, comprising: inserting a stent into an obstructed blood vessel, the stent having a generally tubular structure, the surface of the structure being coated with (or otherwise adapted to release) a composition comprising a compound having the structure:



or pharmaceutically acceptable salt, ester, or salt of ester thereof; wherein  $R_1$  is hydrogen, straight or branched  $C_{1-6}$  heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl;

R<sub>2</sub> and R<sub>3</sub> are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, straight or branched C<sub>1-6</sub>heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl; or

 $R_1$  and  $R_2$ , when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen;

R<sub>4</sub> is hydrogen or halogen;

R<sub>5</sub> is hydrogen or a protecting group;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;

m is 0-2;

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2-R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2-R_{14}$ ;

wherein  $R_{12}$  and  $R_{13}$  are, independently for each occurrence, hydrogen,  $C_{1-6}$ alkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or a protecting group, or  $R_{12}$  and  $R_{13}$ , taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of  $R_{12}$  and  $R_{13}$  are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or -N(alkyl), or wherein  $X_2$ - $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is  $-(C=0)NHR_{15}$   $-(C=0)OR_{15}$ , or  $-(C=0)R_{15}$ , wherein each occurrence of  $R_{15}$  is independently hydrogen, alkyl, heteroalkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or  $R_{14}$  is  $-SO_2(R_{16})$ , wherein  $R_{16}$  is an alkyl moiety, wherein one or more of  $R_{14}$ ,  $R_{15}$ , or  $R_{16}$  are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

 $R_{10}$  is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;  $R_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH2 or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or C<sub>1-6</sub>alkyl, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond; and optionally

a pharmaceutically acceptable carrier or diluent;



such that the obstruction is eliminated and the composition is delivered in amounts effective to prevent or reduce the rate of restenosis;

with the proviso that the following groups do not occur simultaneously as defined: n is 1; X is O;  $R_1$  is methyl;  $R_2$ ,  $R_3$ ,  $R_4$ ,  $R_7$ ,  $R_8$  and  $R_{11}$  are each hydrogen;  $R_5$  is hydrogen,  $C_{1-4}$ alkyl or  $-C(=O)C_{1-4}$ alkyl;  $R_6$  is hydrogen, OH,  $C_{1-4}$ alkoxy or  $-OC(=O)C_{1-4}$ alkyl;  $R_9$  and  $R_{10}$  are independently OH,  $C_{1-4}$ alkoxy or  $-OC(=O)C_{1-4}$ alkyl;

and Y-Z is -CHRYCHRZ-, -CH=CH- or ; wherein RY and RZ are independently hydrogen, C<sub>1-4</sub>alkyl or C<sub>1-4</sub>alkanoyl.

# 124. A method for expanding the lumen of a body passageway, comprising: inserting a stent into the passageway, the stent having a generally tubular structure, the surface of the structure being coated with (or otherwise adapted to release) a composition comprising a compound having the structure:

$$\begin{array}{c|c}
R_{11} & R_{3} & R_{2} \\
R_{11} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} \\
\hline
R_{11} & R_{2} & R_{3} \\
\hline
R_{2} & R_{3} & R_{2} \\
\hline
R_{3} & R_{2} & R_{3} \\
\hline
R_{4} & R_{5} & R_{6} \\
\hline
R_{7} & R_{8} & R_{9} \\
\hline
R_{11} & R_{2} & R_{2} \\
\hline
R_{11} & R_{2} & R_{2} \\
\hline
R_{2} & R_{3} & R_{2} \\
\hline
R_{2} & R_{3} & R_{2} \\
\hline
R_{3} & R_{2} & R_{3} \\
\hline
R_{4} & R_{5} & R_{6} \\
\hline
R_{5} & R_{6} & R_{6} \\
\hline
R_{7} & R_{8} & R_{9} \\
\hline
R_{11} & R_{2} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{12} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{12} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{12} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{12} & R_{2} & R_{3} & R_{2} \\
\hline
R_{12} & R_{2} & R_{3} & R_{2} \\
\hline
R_{11} & R_{2} & R_{3} & R_{2} \\
\hline
R_{12} & R_{2} & R_{3} & R_{2} \\
\hline
R_{12} & R_{2} & R_{3} & R_{3} \\
\hline
R_{12} & R_{2} & R_{3} & R_{3} \\
\hline
R_{12} & R_{2} & R_{3} & R_{3} \\
\hline
R_{12} & R_{2} & R_{3} & R_{3} \\
\hline
R_{12} & R_{2} & R_{3} & R_{3} \\
\hline
R_{12} & R_{2} & R_{3} & R_{3} \\
\hline
R_{12} & R_{2} & R_{3} & R_{3} \\
\hline
R_{13} & R_{2} & R_{3} & R_{3} \\
\hline
R_{11} & R_{2} & R_{3} & R_{3} \\
\hline
R_{12} & R_{3} & R_{3} & R_{3} \\
\hline
R_{13} & R_{2} & R_{3} & R_{3} \\
\hline
R_{13} & R_{2} & R_{3} & R_{3} \\
\hline
R_{13} & R_{2} & R_{3} & R_{3} \\
\hline
R_{13} & R_{3} & R_{3} & R_{3} \\
\hline
R_{13} & R_{3} & R_{3} & R_{3} \\
\hline
R_{11} & R_{3} & R_{3} & R_{3} \\
\hline
R_{12} & R_{3} & R_{3} & R_{3} \\
\hline
R_{13} & R_{3} & R_{3} & R_{3} \\
\hline
R_{11} & R_{3} & R_{3} & R_{3} \\
\hline
R_{12} & R_{3} & R_{3} & R_{3} \\
\hline
R_{13} & R_{3} & R_{3} & R_{3} \\
\hline
R_{11} & R_{3} & R_{3} & R_{3} \\
\hline
R_{12} & R_{3} & R_{3} & R_{3} \\
\hline
R_{12} & R_{3}$$

or pharmaceutically acceptable salt, ester, or salt of ester thereof; wherein  $R_1$  is hydrogen, straight or branched  $C_{1-6}$ alkyl, straight or branched  $C_{1-6}$ heteroallkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl;

 $\mathbb{R}_2$  and  $\mathbb{R}_3$  are each independently hydrogen, halogen, hydroxyl, protected hydroxyl, straight or branched  $\mathbb{C}_{1\text{-}6}$ heteroalkyl, or aryl,

wherein the alkyl, heteroalkyl, and aryl groups may optionally be substituted with one or more occurrences of halogen, hydroxyl or protected hydroxyl; or

R<sub>1</sub> and R<sub>2</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen; or

R<sub>1</sub> and R<sub>3</sub>, when taken together, may form a saturated or unsaturated cyclic ring of 3 to 8 carbon atoms, optionally substituted with one or more occurrences of halogen;

R4 is hydrogen or halogen;

R₅ is hydrogen or a protecting group;

R<sub>6</sub> is hydrogen, hydroxyl, or protected hydroxyl;

n is 0-2;

R<sub>7</sub>, for each occurrence, is independently hydrogen, hydroxyl, or protected hydroxyl;

 $R_8$  is hydrogen, halogen, hydroxyl, protected hydroxyl, alkyloxy, or  $C_{l-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl,  $SR_{12}$ , or  $NR_{12}R_{13}$ ;

 $R_9$  is hydrogen, halogen, hydroxyl, protected hydroxyl,  $OR_{12}$ ,  $SR_{12}$ ,  $NR_{12}R_{13}$ , -  $X_1(CH_2)_pX_2$ - $R_{14}$ , or is  $C_{1-6}$ alkyl optionally substituted with hydroxyl, protected hydroxyl, halogen, amino, protected amino, or - $X_1(CH_2)_pX_2$ - $R_{14}$ ;

wherein R<sub>12</sub> and R<sub>13</sub> are, independently for each occurrence, hydrogen, C<sub>1-6</sub>alkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or a protecting group, or R<sub>12</sub> and R<sub>13</sub>, taken together may form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms, and each of R<sub>12</sub> and R<sub>13</sub> are optionally further substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen,

wherein  $X_1$  and  $X_2$  are each independently absent, or are oxygen, NH, or -N(alkyl), or wherein  $X_2$ - $R_{14}$  together are  $N_3$  or are a saturated or unsaturated heterocyclic moiety,

p is 2-10, and

 $R_{14}$  is hydrogen, or an aryl, heteroaryl, alkylaryl, or alkylheteroaryl moiety, or is  $-(C=O)NHR_{15}$   $-(C=O)OR_{15}$ , or  $-(C=O)R_{15}$ , wherein each



occurrence of  $R_{15}$  is independently hydrogen, alkyl, heteroalkyl, aryl, heteroaryl, alkylaryl, or alkylheteroaryl, or  $R_{14}$  is  $-SO_2(R_{16})$ , wherein  $R_{16}$  is an alkyl moiety, wherein one or more of  $R_{14}$ ,  $R_{15}$ , or  $R_{16}$  are optionally substituted with one or more occurrences of hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen; or

R<sub>8</sub> and R<sub>9</sub> may, when taken together, form a saturated or unsaturated cyclic ring containing 1 to 4 carbon atoms and 1 to 3 nitrogen or oxygen atoms and is optionally substituted with hydroxyl, protected hydroxyl, alkyloxy, amino, protected amino, alkylamino, aminoalkyl, or halogen;

 $R_{10}$  is hydrogen, hydroxyl, protected hydroxyl, amino, or protected amino;  $R_{11}$  is hydrogen, hydroxyl or protected hydroxyl;

X is absent or is O, NH, N-alkyl, CH2 or S;

Y is CHR<sub>17</sub>, O, C=O, CR<sub>17</sub> or NR<sub>17</sub>; and Z is CHR<sub>18</sub>, O, C=O, CR<sub>18</sub> or NR<sub>18</sub>, wherein each occurrence of R<sub>17</sub> and R<sub>18</sub> is independently hydrogen or C<sub>1-6</sub>alkyl, or R<sub>17</sub> and R<sub>18</sub> taken together is -O-,  $-CH_2$ - or  $-NR_{19}$ -, wherein R<sub>19</sub> is hydrogen or C<sub>1-6</sub>alkyl, and Y and Z may be connected by a single or double bond; and optionally

a pharmaceutically acceptable carrier or diluent; such that the passageway is expanded.

- 125. The method of claim 124, wherein the lumen of a body passageway is expanded in order to eliminate a biliary, gastrointestinal, esophageal, tracheal/bronchial, urethral and/or vascular obstruction.
- 126. The method of claim 125, wherein the lumen of a body passageway is expanded in order to eliminate a vascular obstruction.



# This Page is Inserted by IFW Indexing and Scanning Operations and is not part of the Official Record

#### **BEST AVAILABLE IMAGES**

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images include but are not limited to the items checked:

□ BLACK BORDERS
□ IMAGE CUT OFF AT TOP, BOTTOM OR SIDES
□ FADED TEXT OR DRAWING
□ BLURRED OR ILLEGIBLE TEXT OR DRAWING
□ SKEWED/SLANTED IMAGES
□ COLOR OR BLACK AND WHITE PHOTOGRAPHS
□ GRAY SCALE DOCUMENTS
□ LINES OR MARKS ON ORIGINAL DOCUMENT
□ REFERENCE(S) OR EXHIBIT(S) SUBMITTED ARE POOR QUALITY

#### IMAGES ARE BEST AVAILABLE COPY.

OTHER:

As rescanning these documents will not correct the image problems checked, please do not report these problems to the IFW Image Problem Mailbox.